256411

Scientific and Technical Information Center

SEARCH REQUEST FORM

Requester's Full Name: Cecilia Jaisle Examiner #: [redacted] Date: 4-3-08
Art Unit: 1624 Phone Number: 2-4931 Serial Number: 10593590
Location (Bldg/Room#): REM 5B28 (Mailbox #): 5C18 Results Format Preferred (circle): PAPER DISK

To ensure an efficient and quality search, please attach a copy of the cover sheet, claims, and abstract or fill out the following:

Title of Invention: See Bib Data Sheet

Inventors (please provide full names):

Earliest Priority Date:

Search Topic:
Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc., if known.

For Sequence Searches Only Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

See claims attached. Please do structure search and inventor name(s) search. Display results to show identification of source, and RN #, compound name & structure of identified compounds. Search compounds of Formula I where X is CH with additions as noted

Please call with any questions

STAFF USE ONLY

Searcher:	Type of Search	Vendors and cost where applicable
Searcher Phone #:	NA Sequence (#)	STN
Searcher Location:	AA Sequence (#)	Dialog
Date Searcher Picked Up:	Structure (#)	Questel/Orbit
Date Completed:	Bibliographic	Lexis/Nexis
Searcher Prep & Review Time:	Litigation	WWW/Internet
Online Time:	Fulltext	In-house sequence systems
	Other	Commercial Oligomer Score/Length Interference SPDI Encode/Transl Other (specify)

Author Search

```
=> FILE HCAPLUS
FILE 'HCAPLUS' ENTERED AT 14:36:23 ON 11 APR 2008
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FILE COVERS 1907 - 11 Apr 2008  VOL 148 ISS 16
FILE LAST UPDATED: 10 Apr 2008  (20080410/ED)

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 This file contains CAS Registry Numbers for easy and accurate
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'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

=> D QUE L13
L1              STR
```

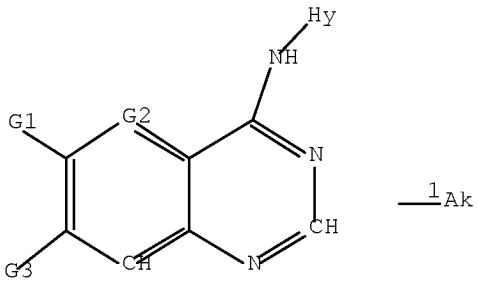


```
G1 O,S
G2 CH,[@1]
G3 H,X,Ak

Structure attributes must be viewed using STN Express query preparation:
Uploading strA.str
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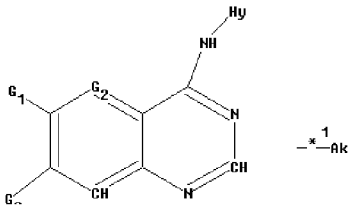

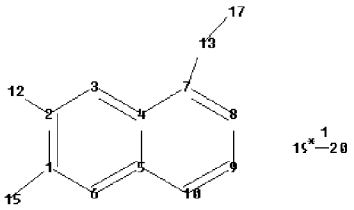

chain nodes :
12 13 15 17 19 20
ring nodes :
1 2 3 4 5 6 7 8 9 10
chain bonds :
1-15 2-12 7-13 13-17 19-20
ring bonds :
1-2 1-6 2-3 3-4 4-5 4-7 5-6 5-10 7-8 8-9 9-10
exact/norm bonds :
1-2 1-6 1-15 2-3 2-12 3-4 4-5 4-7 5-6 5-10 7-8 7-13 8-9 9-10 13-17
19-20

G1:O,S

G2:CH,[*1]

G3:H,X,Ak

Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
12:CLASS 13:CLASS 15:CLASS 17:Atom 19:CLASS 20:CLASS
Generic attributes :
17:
Saturation : Unsaturated

Element Count :
Node 17: Limited
N,N1

L3 150 SEA FILE=REGISTRY SSS FUL L1
L4 6 SEA FILE=HCAPLUS ABB=ON PLU=ON L3

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L5             6 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L4 AND (PRY<=2005 OR AY<=2005
                 OR PY<=2005)
L6           138 SEA FILE=HCAPLUS ABB=ON  PLU=ON  MITSUYA M?/AU
L7            47 SEA FILE=HCAPLUS ABB=ON  PLU=ON  BAMBA M?/AU
L8          7626 SEA FILE=HCAPLUS ABB=ON  PLU=ON  SASAKI Y?/AU
L9          6232 SEA FILE=HCAPLUS ABB=ON  PLU=ON  NISHIMURA T?/AU
L10           19 SEA FILE=HCAPLUS ABB=ON  PLU=ON  EIKI J?/AU
L11         1742 SEA FILE=HCAPLUS ABB=ON  PLU=ON  ARAKAWA K?/AU
L12        15764 SEA FILE=HCAPLUS ABB=ON  PLU=ON  (L6 OR L7 OR L8 OR L9 OR L10
                 OR L11)
L13            1 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L5 AND L12


=> FILE WPIX
FILE 'WPIX' ENTERED AT 14:36:31 ON 11 APR 2008
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FILE LAST UPDATED:                          8 APR 2008   <20080408/UP>
MOST RECENT THOMSON SCIENTIFIC UPDATE:     200823      <200823/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> IPC Reform backfile reclassification has been loaded to the end of
    November 2007. No update date (UP) has been created for the
    reclassified documents, but they can be identified by
    20060101/UPIC and 20061231/UPIC, 20070601/UPIC, 20071001/UPIC and
    20071130/UPIC. <<<

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FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE
http://scientific.thomson.com/support/patents/coverage/latestupdates/

EXPLORE DERWENT WORLD PATENTS INDEX IN STN ANAVIST, VERSION 2.0:
http://www.stn-international.com/archive/presentations/DWPIAnaVist2_0710.pdf

>>> XML document distribution format now available - See HELP XMLDOC <<<

>>> ECLA Codes and Current US National Classifications have been added -
    see NEWS and HELP CHANGE <<<

>>> HELP for European Patent Classifications see HELP ECLA, HELP ICO <<<

>>> Updated PDF files in the following links:
    http://www.stn-international.de/stndatabases/details/ico_0803.zip
    http://www.stn-international.de/stndatabases/details/epc_0803.zip
    Supplement of all changed ECLA items:
    http://www.stn-international.de/stndatabases/details/ecla_0803s.zip <<<
'BI,ABEX' IS DEFAULT SEARCH FIELD FOR 'WPIX' FILE

=> D QUE L17
L1             STR
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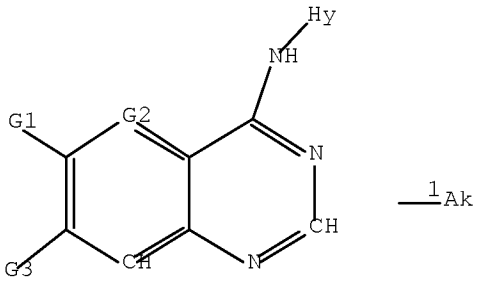

G1 O,S
G2 CH,[@1]
G3 H,X,Ak

Structure attributes must be viewed using STN Express query preparation.

```
L6          138 SEA FILE=HCAPLUS ABB=ON  PLU=ON  MITSUYA M?/AU
L7           47 SEA FILE=HCAPLUS ABB=ON  PLU=ON  BAMBA M?/AU
L8         7626 SEA FILE=HCAPLUS ABB=ON  PLU=ON  SASAKI Y?/AU
L9         6232 SEA FILE=HCAPLUS ABB=ON  PLU=ON  NISHIMURA T?/AU
L10          19 SEA FILE=HCAPLUS ABB=ON  PLU=ON  EIKI J?/AU
L11        1742 SEA FILE=HCAPLUS ABB=ON  PLU=ON  ARAKAWA K?/AU
L12       15764 SEA FILE=HCAPLUS ABB=ON  PLU=ON  (L6 OR L7 OR L8 OR L9 OR L10
                 OR L11)
L15          63 SEA FILE=WPIX SSS FUL L1
L16           2 SEA FILE=WPIX ABB=ON  PLU=ON  L15/DCR
L17           1 SEA FILE=WPIX ABB=ON  PLU=ON  L12 AND L16


=> DUP REM L13 L17
FILE 'HCAPLUS' ENTERED AT 14:36:40 ON 11 APR 2008
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PROCESSING COMPLETED FOR L13
PROCESSING COMPLETED FOR L17
L22           1 DUP REM L13 L17 (1 DUPLICATE REMOVED)
                ANSWER '1' FROM FILE HCAPLUS

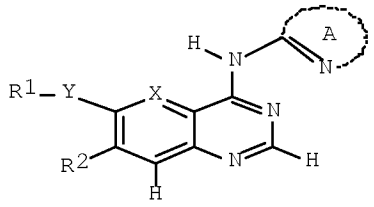
=> D IBIB ED ABS FHITSTR L22 1

L22  ANSWER 1 OF 1  HCAPLUS  COPYRIGHT 2008 ACS on STN DUPLICATE 1
ACCESSION NUMBER:         2005:1042235  HCAPLUS  Full-text
DOCUMENT NUMBER:          143:347192
TITLE:                    Preparation of substituted quinazoline and
                          pyridopyrimidine derivatives as glucokinase activators
INVENTOR(S):              Mitsuya, Morihiro; Bamba, Makoto;
                          Sasaki, Yasuhiro; Nishimura, Teruyuki
                          ; Eiki, Junichi; Arakawa, Keisuke
PATENT ASSIGNEE(S):       Banyu Pharmaceutical Co., Ltd, Japan
SOURCE:                   PCT Int. Appl., 192 pp.
                          CODEN: PIXXD2
DOCUMENT TYPE:            Patent
LANGUAGE:                 Japanese
```

FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005090332	A1	20050929	WO 2005-JP5991	20050323 <--
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2005223610	A1	20050929	AU 2005-223610	20050323 <--
CA 2560286	A1	20050929	CA 2005-2560286	20050323 <--
EP 1734040	A1	20061220	EP 2005-721640	20050323 <--
R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, LV				
CN 1934100	A	20070321	CN 2005-80009447	20050323 <--
IN 2006DN05151	A	20070803	IN 2006-DN5151	20060907 <--
US 20080032996	A1	20080207	US 2007-593540	20070510 <--
PRIORITY APPLN. INFO.:			JP 2004-85808 A	20040323 <--
			WO 2005-JP5991 W	20050323 <--

OTHER SOURCE(S): MARPAT 143:347192
ED Entered STN: 29 Sep 2005
GI

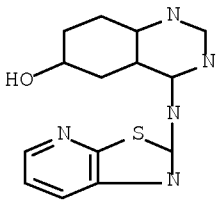


I

AB The title compds. I [X is a nitrogen atom, CH; Y is O, S; R1 is an optionally substituted 5 to 6-membered heteroaryl group, aryl, alkyl, etc.; R2 is a hydrogen atom or a fluorine atom; and the ring A is an optionally substituted monocyclic or bicyclic heteroaryl group] are prepared Thus, [6-(4H-[1,2,4]triazol-3-ylsulfanyl)quinazolin-4-yl]thiazolo[5,4- b]pyridin-2-ylamine was prepared in 2 steps from 4-chloro-6-iodoquinazoline and thiazolo[5,4-b]pyridin-2-ylamine. In a test for glucokinase activating activity, compds. of this invention showed EC50 values of 0.08 μM to 0.18 μM. Formulations are given.

IT 865662-62-0P
RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
(preparation of substituted quinazoline and pyridopyrimidine derivs. as glucokinase activators)

RN 865662-62-0 HCAPLUS
CN 6-Quinazolinol, 4-(thiazolo[5,4-b]pyridin-2-ylamino)- (CA INDEX NAME)



ONE OR MORE TAUTOMERIC DOUBLE BONDS NOT DISPLAYED IN THE STRUCTURE
REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Structure Search

```
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FILE COVERS 1907 - 11 Apr 2008  VOL 148 ISS 16
FILE LAST UPDATED: 10 Apr 2008  (20080410/ED)

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=> D QUE L5
L1              STR
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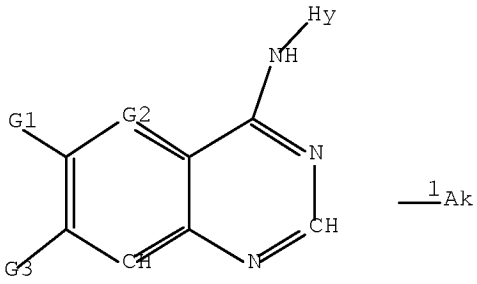


```
G1 O,S
G2 CH,[@1]
G3 H,X,Ak

Structure attributes must be viewed using STN Express query preparation.
L3          150 SEA FILE=REGISTRY SSS FUL L1
L4            6 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L3
L5            6 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L4 AND (PRY<=2005 OR AY<=2005
                OR PY<=2005)


=> S L5 NOT L13
L23           5 L5 NOT L13
```

```
=> FILE WPIX
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FILE LAST UPDATED:                          8 APR 2008   <20080408/UP>
MOST RECENT THOMSON SCIENTIFIC UPDATE:     200823      <200823/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> IPC Reform backfile reclassification has been loaded to the end of
    November 2007. No update date (UP) has been created for the
    reclassified documents, but they can be identified by
    20060101/UPIC and 20061231/UPIC, 20070601/UPIC, 20071001/UPIC and
    20071130/UPIC. <<<

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http://scientific.thomson.com/support/patents/coverage/latestupdates/

EXPLORE DERWENT WORLD PATENTS INDEX IN STN ANAVIST, VERSION 2.0:
http://www.stn-international.com/archive/presentations/DWPIAnaVist2_0710.pdf

>>> XML document distribution format now available - See HELP XMLDOC <<<

>>> ECLA Codes and Current US National Classifications have been added -
    see NEWS and HELP CHANGE <<<

>>> HELP for European Patent Classifications see HELP ECLA, HELP ICO <<<

>>> Updated PDF files in the following links:
    http://www.stn-international.de/stndatabases/details/ico_0803.zip
    http://www.stn-international.de/stndatabases/details/epc_0803.zip
    Supplement of all changed ECLA items:
    http://www.stn-international.de/stndatabases/details/ecla_0803s.zip <<<
'BI,ABEX' IS DEFAULT SEARCH FIELD FOR 'WPIX' FILE

=> D QUE L16
L1              STR
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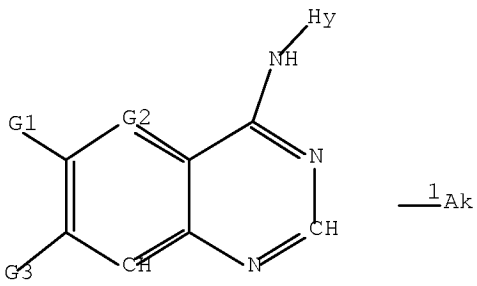


```
G1 O,S
G2 CH,[@1]
G3 H,X,Ak
```

Structure attributes must be viewed using STN Express query preparation.

```
L15          63 SEA FILE=WPIX SSS FUL L1
L16           2 SEA FILE=WPIX ABB=ON  PLU=ON  L15/DCR


=> S L16 NOT L17
L24          1 L16 NOT L17

=> FILE BABS
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FILE LAST UPDATED: 17 MAR 2008       <20080317/UP>
FILE COVERS 1980 TO DATE.

=> D QUE L20
L20           1 SEA FILE=BABS ABB=ON  PLU=ON  6424720/BABSAN


=> FILE BEILSTEIN
FILE 'BEILSTEIN' ENTERED AT 14:37:44 ON 11 APR 2008
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FILE LAST UPDATED ON April 1, 2008

FILE COVERS 1771 TO 2008.
*** FILE CONTAINS 10.322,808  SUBSTANCES ***

>>>PLEASE NOTE: Reaction Data and substance data are stored in
   separate documents and can not be searched together in one query.
   Reaction data for BEILSTEIN compounds may be displayed
   immediately with the display codes PRE (preparations) and REA
   (reactions).  A substance answer set retrieved after the search
   for a chemical name, a compounds with available reaction
   information by combining with PRE/FA, REA/FA or more generally
   with RX/FA. The BEILSTEIN Registry Number (BRN) is the link
   between a BEILSTEIN compound and belonging reactions. For mo
   detailed reaction searches BRNs can be searched as reaction
   partner BRNs Reactant BRN (RX.RBRN) or Product BRN (RX.PBRN).<<<

>>> FOR SEARCHING PREPARATIONS SEE HELP PRE  <<<

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**********************************************************************

>>> Price change as of January 1st, 2008: Connect Time and Structure
    Search fees re-introduced. See NEWS and HELP COST <<<

=> D QUE L21
L1             STR
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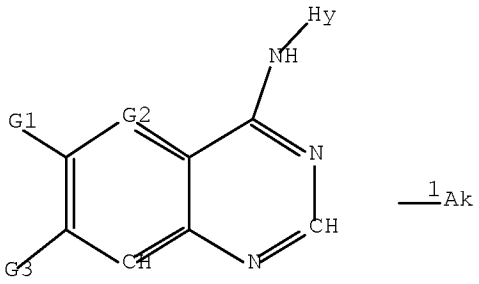

G1 O,S
G2 CH,[@1]
G3 H,X,Ak

Structure attributes must be viewed using STN Express query preparation.
L18 4 SEA FILE=BEILSTEIN SSS FUL L1
L19 1 SEA FILE=BEILSTEIN ABB=ON PLU=ON L18 AND BABSAN/FA
L21 3 SEA FILE=BEILSTEIN ABB=ON PLU=ON L18 NOT L19

=> DUP REM L23 L24 L20 L21
DUPLICATE IS NOT AVAILABLE IN 'BEILSTEIN'.
ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE
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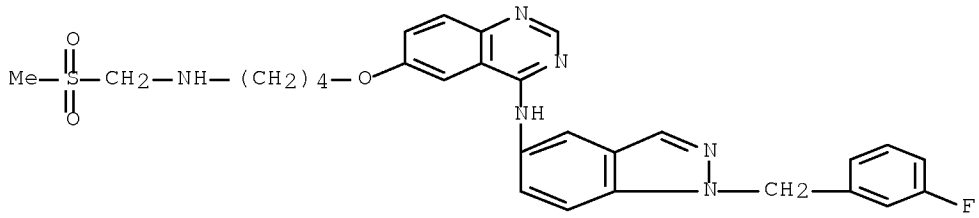
PROCESSING COMPLETED FOR L23
PROCESSING COMPLETED FOR L24
PROCESSING COMPLETED FOR L20
PROCESSING COMPLETED FOR L21
L25 8 DUP REM L23 L24 L20 L21 (2 DUPLICATES REMOVED)
ANSWERS '1-5' FROM FILE HCAPLUS
ANSWERS '6-8' FROM FILE BEILSTEIN

=> D IBIB ED ABS HITSTR 1-5; D IDE ALLREF 6-8

L25 ANSWER 1 OF 8 HCAPLUS COPYRIGHT 2008 ACS on STN DUPLICATE 1
ACCESSION NUMBER: 2003:981459 HCAPLUS Full-text
DOCUMENT NUMBER: 140:228690
TITLE: Synthesis and SAR of potent EGFR/erbB2 dual inhibitors
AUTHOR(S): Zhang, Yue-Mei; Cockerill, Stuart; Guntrip, Stephen B.; Rusnak, David; Smith, Kathryn; Vanderwall, Dana; Wood, Edgar; Lackey, Karen
CORPORATE SOURCE: GlaxoSmithKline, Research Triangle Park, NC, 27709,

USA
SOURCE: Bioorganic & Medicinal Chemistry Letters (2004), 14(1), 111-114
CODEN: BMCLE8; ISSN: 0960-894X
PUBLISHER: Elsevier Science B.V.
DOCUMENT TYPE: Journal
LANGUAGE: English
ED Entered STN: 17 Dec 2003
AB A series of 6-alkoxy-4-anilinoquinazoline compds. was prepared and evaluated for in vitro inhibition of the erbB2 and EGFR kinase activity. The IC50 values of the best compds. were below 0.10 uM. Further, several of these compds. inhibit the growth of erbB2 and EGFR over-expressing tumor cell lines at concns. below 1 uM.
IT 668437-13-6P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(synthesis and structure-activity relations of potent EGFR/erbB2 kinase dual inhibitors)
RN 668437-13-6 HCAPLUS
CN 4-Quinazolinamine, N-[1-[(3-fluorophenyl)methyl]-1H-indazol-5-yl]-6-[4-[[(methylsulfonyl)methyl]amino]butoxy]- (CA INDEX NAME)



REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

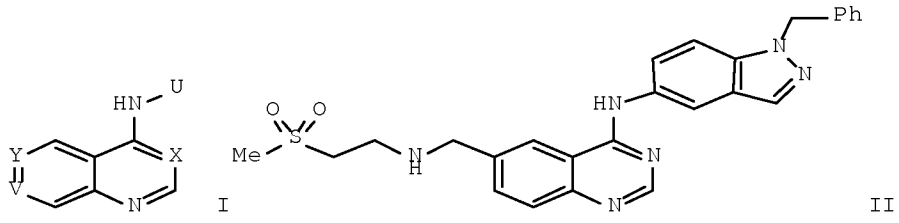
L25 ANSWER 2 OF 8 HCAPLUS COPYRIGHT 2008 ACS on STN DUPLICATE 2
ACCESSION NUMBER: 1999:451283 HCAPLUS Full-text
DOCUMENT NUMBER: 131:102287
TITLE: Preparation of quinazolinylamines and analogs as protein tyrosine kinase inhibitors
INVENTOR(S): Cockerill, George Stuart; Lackey, Karen Elizabeth
PATENT ASSIGNEE(S): Glaxo Group Limited, UK
SOURCE: PCT Int. Appl., 145 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9935132	A1	19990715	WO 1999-GB76	19990111 <--

W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,

TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

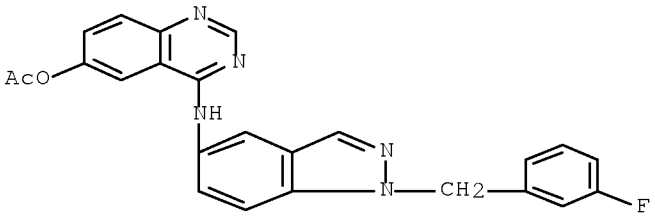
AU 9919786 A 19990726 AU 1999-19786 19990111 <--
PRIORITY APPLN. INFO.: GB 1998-575 A 19980112 <--
WO 1999-GB76 W 19990111 <--
OTHER SOURCE(S): MARPAT 131:102287
ED Entered STN: 23 Jul 1999
GI



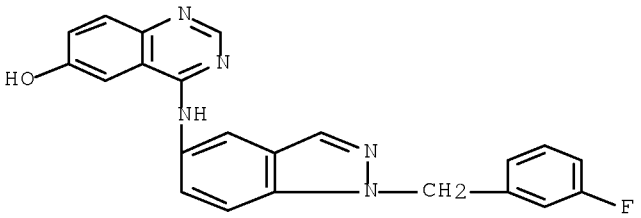
AB Substituted heteroarom. compds. I are prepared [wherein X = N or CH; Y = CR1 and V = N; or Y = N and V = CR1; or Y = CR1 and V = CR2; or Y = CR2 and V = CR1; R1 = Q-M-, wherein M = C1-5 alkylene where any C atom not immediately adjacent to Q may be replaced by O, S, or NR6; Q = wide variety of groups; R2 = H, halo, OH, alkyl, alkoxy, (di)alkylamino; U = Ph, pyridyl, pyrimidinyl, imidazolyl, or 9- or 10-membered bicyclic heterocyclyl containing 1-2 N atoms and 0-1 addnl. O, N, or S; U is substituted by R3, where R3 = benzyl, halobenzyl, pyridylmethyl, pyridylmethoxy, PhO, PhSO2, (un)substituted phthalimido; R6 = H, alkyl]. Twelve examples and a variety of intermediates were prepared For instance, 4-chloro-6-iodoquinazoline was aminated in the 4-position with 5-amino-1-benzyl-1H-indazole, followed by Pd-catalyzed carbonylation, to give 4-[(1-benzyl-1H-indazol-5-yl)amino]quinazoline-6-carbaldehyde. This underwent reductive amination by MeSO2CH2CH2NH2 and a reducing agent such as NaBH(OAc)3, to give title compound II.HCl. In an EGFr phosphorylation assay, II.HCl had an IC50 of <0.10 μM.

IT 230955-59-6P 230955-60-9P 230955-73-4P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; preparation of quinazolinylamines and analogs as protein tyrosine kinase inhibitors)

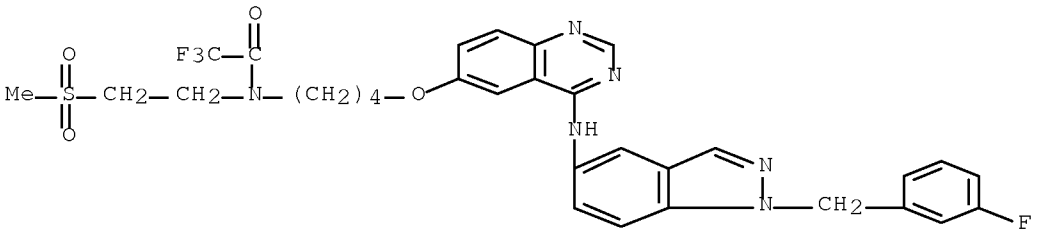
RN 230955-59-6 HCAPLUS
CN 6-Quinazolinol, 4-[[1-[(3-fluorophenyl)methyl]-1H-indazol-5-yl]amino]-, acetate (ester) (9CI) (CA INDEX NAME)



RN 230955-60-9 HCAPLUS
CN 6-Quinazolinol, 4-[[1-[(3-fluorophenyl)methyl]-1H-indazol-5-yl]amino]-
(CA INDEX NAME)

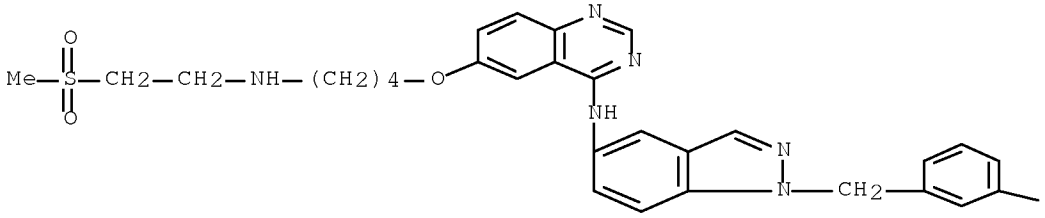


RN 230955-73-4 HCAPLUS
CN Acetamide, 2,2,2-trifluoro-N-[4-[[4-[[1-[(3-fluorophenyl)methyl]-1H-
indazol-5-yl]amino]-6-quinazolinyl]oxy]butyl]-N-[2-(methylsulfonyl)ethyl]-
(CA INDEX NAME)



IT 230955-49-4P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(target compound; preparation of quinazolinylamines and analogs as protein tyrosine kinase inhibitors)
RN 230955-49-4 HCAPLUS
CN 4-Quinazolinamine, N-[1-[(3-fluorophenyl)methyl]-1H-indazol-5-yl]-6-[4-[[2-(methylsulfonyl)ethyl]amino]butoxy]- (CA INDEX NAME)

PAGE 1-A



PAGE 1-B

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 3 OF 8 HCAPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 2001:743253 HCAPLUS Full-text
DOCUMENT NUMBER: 136:79264
TITLE: The characterization of novel, dual ErbB-2/EGFR, tyrosine kinase inhibitors: potential therapy for cancer
AUTHOR(S): Rusnak, David W.; Affleck, Karen; Cockerill, Stuart G.; Stubberfield, Colin; Harris, Robert; Page, Martin; Smith, Kathryn J.; Guntrip, Stephen B.; Carter, Malcolm C.; Shaw, Robert J.; Jowett, Amanda; Stables, Jeremy; Topley, Peter; Wood, Edgar R.; Brignola, Perry S.; Kadwell, Sue H.; Reep, Bryan R.; Mullin, Robert J.; Alligood, Krystal J.; Keith, Barry R.; Crosby, Renae M.; Murray, Doris M.; Knight, W. Blaine; Gilmer, Tona M.; Lackey, Karen
CORPORATE SOURCE: Department of Cancer Biology, GlaxoSmithKline, Research Triangle Park, NC, 27709, USA
SOURCE: Cancer Research (2001), 61(19), 7196-7203
CODEN: CNREA8; ISSN: 0008-5472
PUBLISHER: American Association for Cancer Research
DOCUMENT TYPE: Journal
LANGUAGE: English
ED Entered STN: 11 Oct 2001
AB The type 1 receptor tyrosine kinases constitute a family of transmembrane proteins involved in various aspects of cell growth and survival and have been implicated in the initiation and progression of several types of human malignancies. The best characterized of these proteins are the epidermal growth factor receptor (EGFR) and ErbB-2 (HER-2/neu). We have developed potent quinazoline and pyrido-[3,4-d]-pyrimidine small mols. that are dual inhibitors of ErbB-2 and EGFR. The compds. demonstrate potent in vitro inhibition of the ErbB-2 and EGFR kinase domains with IC50s <80 nM. Growth of

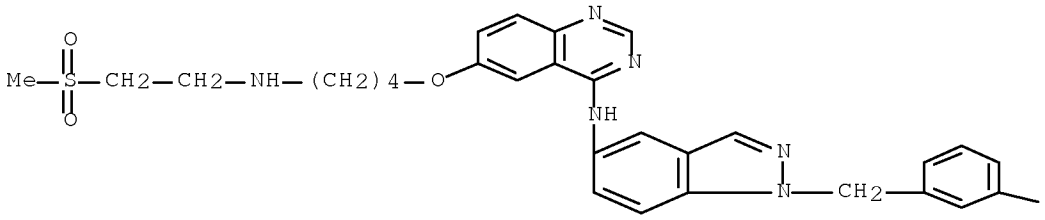
ErbB-2- and EGFR-expressing tumor cell lines is inhibited at concns. <0.5 μM. Selectivity for tumor cell growth inhibition vs. normal human fibroblast growth inhibition ranges from 10- to >75-fold. Tumor growth in mouse s.c. xenograft models of the BT474 and HN5 cell lines is inhibited in a dose-responsive manner using oral doses of 10 and 30 mg/kg twice per day. In addition, the tested compds. caused a reduction of ErbB-2 and EGFR autophosphorylation in tumor fragments from these xenograft models. These data indicate that these compds. have potential use as therapy in the broad population of cancer patients overexpressing ErbB-2 and/or EGFR.

IT 230955-49-4, GW 5945
RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(characterization of novel, dual ErbB-2/EGFR, tyrosine kinase inhibitors and potential therapy for cancer)

RN 230955-49-4 HCAPLUS

CN 4-Quinazolinamine, N-[1-[(3-fluorophenyl)methyl]-1H-indazol-5-yl]-6-[4-[[2-(methylsulfonyl)ethyl]amino]butoxy]- (CA INDEX NAME)

PAGE 1-A



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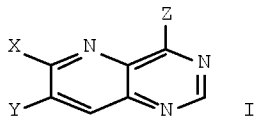
REFERENCE COUNT: 48 THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 4 OF 8 HCAPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 1999:113672 HCAPLUS Full-text
DOCUMENT NUMBER: 130:182476
TITLE: Preparation of heterocyclic compounds as irreversible bicyclic inhibitors of tyrosine kinases
INVENTOR(S): Bridges, Alexander James
PATENT ASSIGNEE(S): Warner-Lambert Company, USA
SOURCE: PCT Int. Appl., 131 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9906396	A1	19990211	WO 1998-US15592	19980729 <--
W: AL, AU, BA, BB, BG, BR, CA, CN, CZ, EE, GE, HR, HU, ID, IL, IS, JP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 9886659	A	19990222	AU 1998-86659	19980729 <--
US 6153617	A	20001128	US 1999-269647	19990325 <--
US 20030087881	A1	20030508	US 2002-272651	20021017 <--
PRIORITY APPLN. INFO.:			US 1997-54061P	P 19970729 <--
			WO 1998-US15592	W 19980729 <--
			US 1999-269647	A3 19990325 <--
			US 2000-656331	B1 20000906 <--

OTHER SOURCE(S): MARPAT 130:182476
ED Entered STN: 19 Feb 1999
GI

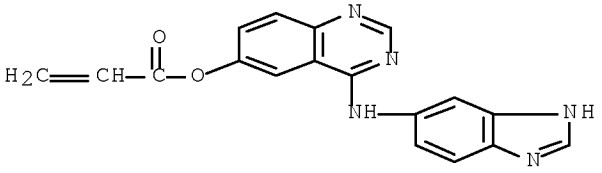


I

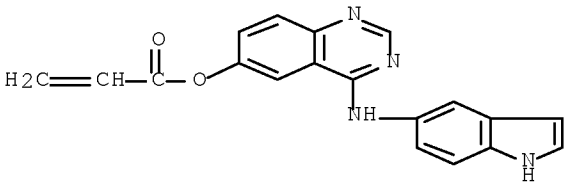
AB The title compds., e.g. I [X = DEF, Y = SR4, etc. ; or X = SR4, etc., and Y = DEF; D = O, etc.; E = CO, etc.; F = CR1(:C):C(R5)H, etc.; a proviso is given; R1 = H, halo, etc.; R5 = H, halo, perfluoroalkyl, etc.; Z = indoline moiety (generic structure given), etc.; R4 = H, alkyl, etc.], are prepared This invention also provides a method of treating cancer, restenosis, atherosclerosis, endometriosis, and psoriasis and a pharmaceutical composition that comprises a compound that is an irreversible inhibitor of tyrosine kinases. N-[4-(6-bromo-2,3-dihydroindol-1- yl)quinazolin-6-yl]acrylamide in vitro showed IC50 of 0.4 nM against epidermal growth factor receptor tyrosine kinase.

IT 220577-65-1P 220577-66-2P 220577-73-1P
220577-74-2P 220577-75-3P 220577-76-4P
220577-77-5P 220577-78-6P 220577-79-7P
220577-80-0P 220577-82-2P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of heterocyclic compds. as irreversible bicyclic inhibitors of tyrosine kinases)

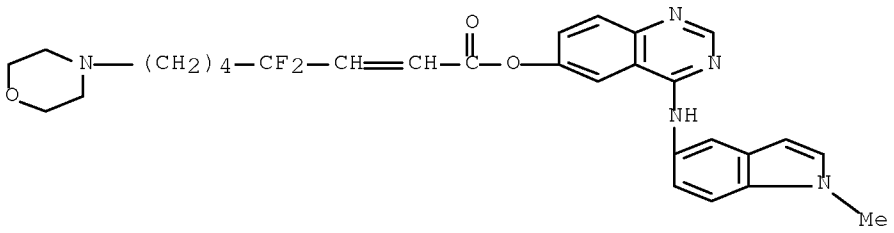
RN 220577-65-1 HCAPLUS
CN 2-Propenoic acid, 4-(1H-benzimidazol-5-ylamino)-6-quinazolinyl ester (9CI) (CA INDEX NAME)



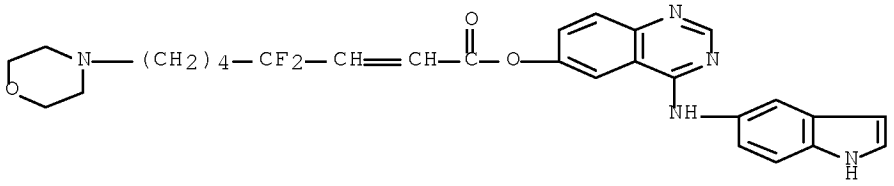
RN 220577-66-2 HCAPLUS
CN 2-Propenoic acid, 4-(1H-indol-5-ylamino)-6-quinazolinyl ester (CA INDEX NAME)



RN 220577-73-1 HCAPLUS
CN 2-Octenoic acid, 4,4-difluoro-8-(4-morpholinyl)-, 4-[(1-methyl-1H-indol-5-yl)amino]-6-quinazolinyl ester (CA INDEX NAME)

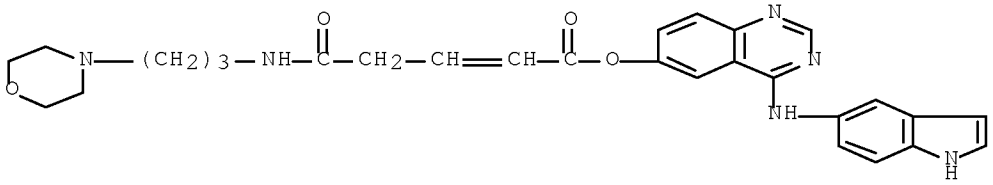


RN 220577-74-2 HCAPLUS
CN 2-Octenoic acid, 4,4-difluoro-8-(4-morpholinyl)-, 4-(1H-indol-5-ylamino)-6-quinazolinyl ester (CA INDEX NAME)



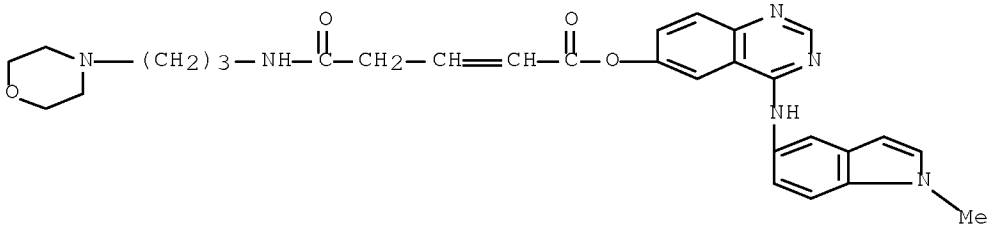
RN 220577-75-3 HCAPLUS

CN 2-Pentenoic acid, 5-[[3-(4-morpholinyl)propyl]amino]-5-oxo-, 4-(1H-indol-5-ylamino)-6-quinazolinyl ester (CA INDEX NAME)



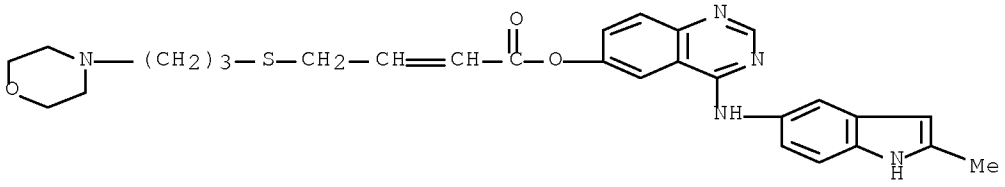
RN 220577-76-4 HCAPLUS

CN 2-Pentenoic acid, 5-[[3-(4-morpholinyl)propyl]amino]-5-oxo-, 4-[(1-methyl-1H-indol-5-yl)amino]-6-quinazolinyl ester (CA INDEX NAME)



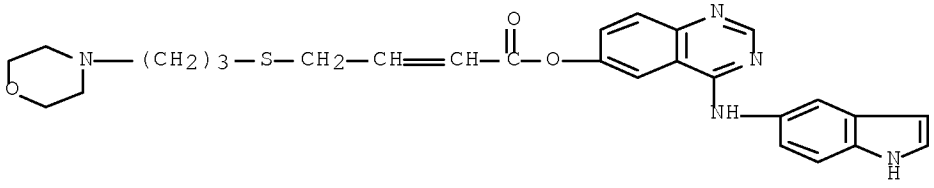
RN 220577-77-5 HCAPLUS

CN 2-Butenoic acid, 4-[[3-(4-morpholinyl)propyl]thio]-, 4-[(2-methyl-1H-indol-5-yl)amino]-6-quinazolinyl ester (CA INDEX NAME)

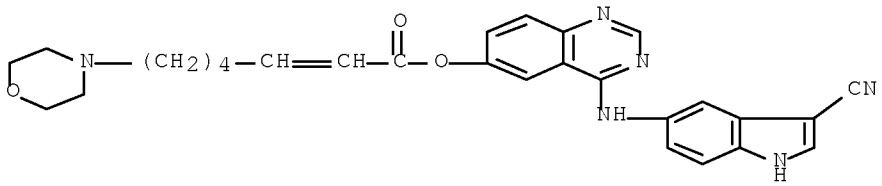


RN 220577-78-6 HCAPLUS

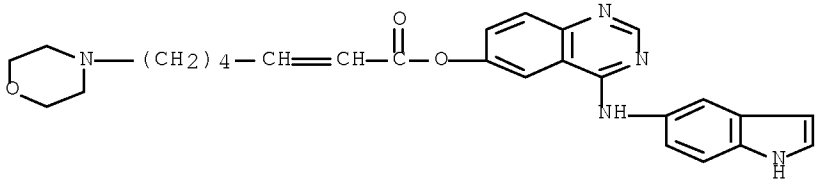
CN 2-Butenoic acid, 4-[[3-(4-morpholinyl)propyl]thio]-, 4-(1H-indol-5-ylamino)-6-quinazolinyl ester (CA INDEX NAME)



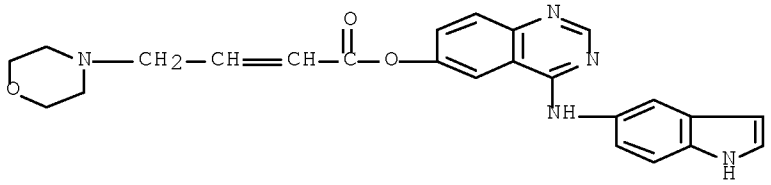
RN 220577-79-7 HCAPLUS
CN 2-Heptenoic acid, 7-(4-morpholinyl)-, 4-[(3-cyano-1H-indol-5-yl)amino]-6-quinazolinyl ester (CA INDEX NAME)



RN 220577-80-0 HCAPLUS
CN 2-Heptenoic acid, 7-(4-morpholinyl)-, 4-(1H-indol-5-ylamino)-6-quinazolinyl ester (CA INDEX NAME)



RN 220577-82-2 HCAPLUS
CN 2-Butenoic acid, 4-(4-morpholinyl)-, 4-(1H-indol-5-ylamino)-6-quinazolinyl ester (CA INDEX NAME)



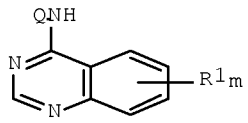
REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 5 OF 8 HCAPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 1995:23238 HCAPLUS Full-text
DOCUMENT NUMBER: 122:31545
TITLE: Preparation of aminoquinazolines useful in the treatment of cancer
INVENTOR(S): Barker, Andrew John; Brown, Dearg Sutherland
PATENT ASSIGNEE(S): Zeneca, UK
SOURCE: Eur. Pat. Appl., 39 pp.

CODEN: EPXXDW
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 602851	A1	19940622	EP 1993-309680	19931203 <--
EP 602851	B1	19961009		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
AU 9350728	A	19940623	AU 1993-50728	19931116 <--
AU 664496	B2	19951116		
ZA 9308594	A	19940610	ZA 1993-8594	19931117 <--
CA 2103383	A1	19940611	CA 1993-2103383	19931118 <--
CA 2103383	C	20050125		
IL 107678	A	19990312	IL 1993-107678	19931119 <--
HU 65622	A2	19940728	HU 1993-3328	19931124 <--
FI 9305431	A	19940611	FI 1993-5431	19931203 <--
AT 143956	T	19961015	AT 1993-309680	19931203 <--
ES 2093367	T3	19961216	ES 1993-309680	19931203 <--
CZ 283612	B6	19980513	CZ 1993-2651	19931206 <--
NO 9304504	A	19940613	NO 1993-4504	19931209 <--
JP 06336481	A	19941206	JP 1993-309184	19931209 <--
JP 3330706	B2	20020930		
CN 1094043	A	19941026	CN 1993-120872	19931210 <--
US 5580870	A	19961203	US 1993-164725	19931210 <--
PRIORITY APPLN. INFO.:			GB 1992-25765 A	19921210 <--
			GB 1993-10248 A	19930518 <--

OTHER SOURCE(S): MARPAT 122:31545
ED Entered STN: 08 Nov 1994
GI

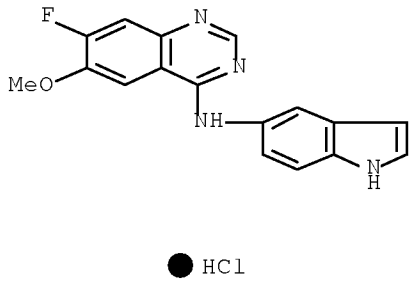


I

AB The title compds. [I; Q = 9- or 10-membered bicyclic heterocyclic moiety containing 1-2 N atoms; R1 = OH, NH2, ureido, hydroxyamino, trifluoromethoxy, (un)substituted C1-4 alkyl, C1-4 alkoxy, pyrrolidin-1-yl, piperidino, etc.; m = 1-3], useful in the treatment of cancer (no data), are prepared and I-containing formulations presented. Thus, 4-chloro-6,7- dimethoxyquinazoline was reacted with 5-aminoquinoline, producing 6,7-dimethoxy-4-(5-quinolylamino)quinazoline, m.p. > 240°, in 35% yield.

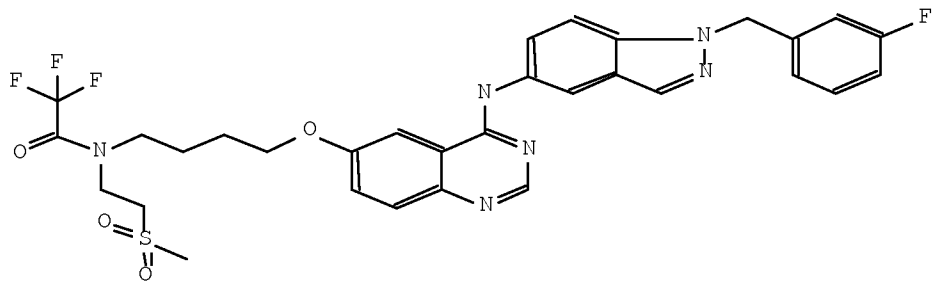
IT 159768-49-7P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of, as anticancer agent)

RN 159768-49-7 HCAPLUS
CN 4-Quinazolinamine, 7-fluoro-N-1H-indol-5-yl-6-methoxy-, monohydrochloride (9CI) (CA INDEX NAME)



L25 ANSWER 6 OF 8 BEILSTEIN COPYRIGHT 2008 BEILSTEIN MDL on STN

Beilstein Records (BRN):	9602965
Chemical Name (CN):	2,2,2-trifluoro-N-(4-<4-<1-(3-fluoro-benzyl)-1H-indazol-5-ylamino>-quinazolin-6-yloxy>-butyl)-N-(2-methanesulfonyl-ethyl)-acetamide
Autonom Name (AUN):	2,2,2-trifluoro-N-(4-<4-<1-(3-fluoro-benzyl)-1H-indazol-5-ylamino>-quinazolin-6-yloxy>-butyl)-N-(2-methanesulfonyl-ethyl)-acetamide
Molec. Formula (MF):	C31 H30 F4 N6 O4 S
Molecular Weight (MW):	658.67
Lawson Number (LN):	29684, 29566, 16445, 3140, 3125, 1157, 292
Compound Type (CTYPE):	heterocyclic
Constitution ID (CONSID):	8097768
Tautomer ID (TAUTID):	9008192
Entry Date (DED):	2004/04/23
Update Date (DUPD):	2004/04/23



Field Availability:

Code	Name	Occurrence

BRN	Beilstein Records	1
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AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	7
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1

This substance also occurs in Reaction Documents:

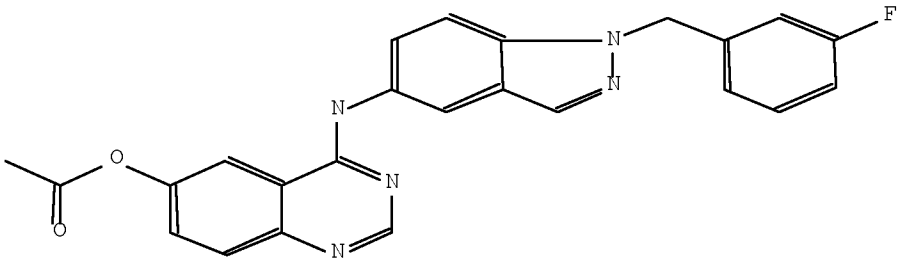
Code	Name	Occurrence
RX	Reaction Documents	2
RXREA	Substance is Reaction Reactant	1
RXPRO	Substance is Reaction Product	1

All References:
ALLREF

1. Zhang, Yue-Mei; Cockerill, Stuart; Guntrip, Stephen B.; Rusnak, David; Smith, Kathryn; Vanderwall, Dana; Wood, Edgar; Lackey, Karen, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 14(1), <2004>, 111 - 114; BABS-6424720

L25 ANSWER 7 OF 8 BEILSTEIN COPYRIGHT 2008 BEILSTEIN MDL on STN

Beilstein Records (BRN):	9589052
Chemical Name (CN):	acetic acid 4-<1-(3-fluoro-benzyl)-1H-indazol-5-ylamino>-quinazolin-6-yl ester
Autonom Name (AUN):	acetic acid 4-<1-(3-fluoro-benzyl)-1H-indazol-5-ylamino>-quinazolin-6-yl ester
Molec. Formula (MF):	C24 H18 F N5 O2
Molecular Weight (MW):	427.44
Lawson Number (LN):	29684, 29566, 16445, 1155
Compound Type (CTYPE):	heterocyclic
Constitution ID (CONSID):	8086336
Tautomer ID (TAUTID):	9002439
Entry Date (DED):	2004/04/23
Update Date (DUPD):	2004/04/23



Field Availability:

Code	Name	Occurrence
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	4
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
RX	Reaction Documents	2
RXREA	Substance is Reaction Reactant	1
RXPRO	Substance is Reaction Product	1

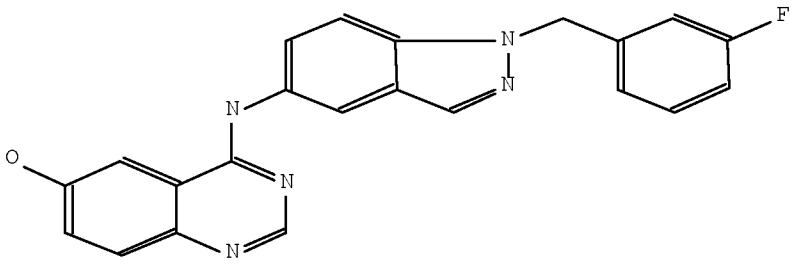
All References:
ALLREF

1. Zhang, Yue-Mei; Cockerill, Stuart; Guntrip, Stephen B.; Rusnak, David; Smith, Kathryn; Vanderwall, Dana; Wood, Edgar; Lackey, Karen, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 14(1), <2004>, 111 - 114; BABS-6424720

L25 ANSWER 8 OF 8 BEILSTEIN COPYRIGHT 2008 BEILSTEIN MDL on STN

Beilstein Records (BRN):	9584302
Chemical Name (CN):	4-<1-(3-fluoro-benzyl)-1H-indazol-5-ylamino>-quinazolin-6-ol
Autonom Name (AUN):	4-<1-(3-fluoro-benzyl)-1H-indazol-5-ylamino>-quinazolin-6-ol
Molec. Formula (MF):	C22 H16 F N5 O
Molecular Weight (MW):	385.40
Lawson Number (LN):	29684, 29566, 16445
Compound Type (CTYPE):	heterocyclic
Constitution ID (CONSID):	8082402

Tautomer ID (TAUTID): 8997062
Entry Date (DED): 2004/04/23
Update Date (DUPD): 2004/04/23



Field Availability:

Code	Name	Occurrence
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	3
FS	File Segment	1
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
RX	Reaction Documents	2
RXREA	Substance is Reaction Reactant	1
RXPRO	Substance is Reaction Product	1

All References:
ALLREF

1. Zhang, Yue-Mei; Cockerill, Stuart; Guntrip, Stephen B.; Rusnak, David; Smith, Kathryn; Vanderwall, Dana; Wood, Edgar; Lackey, Karen, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 14(1), <2004>, 111 - 114; BABS-6424720

Search History

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L1              STRUCTURE UPLOADED
L2           12 SEA SSS SAM L1
L3          150 SEA SSS FUL L1

     FILE 'HCAPLUS' ENTERED AT 14:26:51 ON 11 APR 2008
L4            6 SEA ABB=ON  PLU=ON  L3
L5            6 SEA ABB=ON  PLU=ON  L4 AND (PRY<=2005 OR AY<=2005 OR PY<=2005)

L6          138 SEA ABB=ON  PLU=ON  MITSUYA M?/AU
L7           47 SEA ABB=ON  PLU=ON  BAMBA M?/AU
L8         7626 SEA ABB=ON  PLU=ON  SASAKI Y?/AU
L9         6232 SEA ABB=ON  PLU=ON  NISHIMURA T?/AU
L10          19 SEA ABB=ON  PLU=ON  EIKI J?/AU
L11        1742 SEA ABB=ON  PLU=ON  ARAKAWA K?/AU
L12       15764 SEA ABB=ON  PLU=ON  (L6 OR L7 OR L8 OR L9 OR L10 OR L11)
L13           1 SEA ABB=ON  PLU=ON  L5 AND L12

     FILE 'WPIX' ENTERED AT 14:29:28 ON 11 APR 2008
L14           8 SEA SSS SAM L1
L15          63 SEA SSS FUL L1
L16           2 SEA ABB=ON  PLU=ON  L15/DCR
L17           1 SEA ABB=ON  PLU=ON  L12 AND L16

     FILE 'BEILSTEIN' ENTERED AT 14:33:40 ON 11 APR 2008
L18           4 SEA SSS FUL L1
                SEL BABSAN
L19           1 SEA ABB=ON  PLU=ON  L18 AND BABSAN/FA
                SEL BABSAN

     FILE 'BABS' ENTERED AT 14:34:55 ON 11 APR 2008
L20           1 SEA ABB=ON  PLU=ON  6424720/BABSAN

     FILE 'BEILSTEIN' ENTERED AT 14:35:08 ON 11 APR 2008
L21           3 SEA ABB=ON  PLU=ON  L18 NOT L19

     FILE 'HCAPLUS, WPIX' ENTERED AT 14:36:40 ON 11 APR 2008
L22           1 DUP REM L13 L17 (1 DUPLICATE REMOVED)

     FILE 'HCAPLUS' ENTERED AT 14:36:58 ON 11 APR 2008
                D QUE L5
L23           5 SEA ABB=ON  PLU=ON  L5 NOT L13

     FILE 'WPIX' ENTERED AT 14:37:14 ON 11 APR 2008
L24           1 SEA ABB=ON  PLU=ON  L16 NOT L17

     FILE 'HCAPLUS, WPIX, BABS, BEILSTEIN' ENTERED AT 14:37:59 ON 11 APR 2008
L25           8 DUP REM L23 L24 L20 L21 (2 DUPLICATES REMOVED)
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